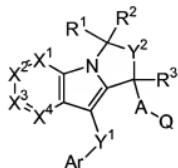


Amendment to the Claims:

Please amend Claims 1, 6, 15 and 20, and cancel Claims 3, 7 and 22-26 as follows.

Listing of Claims:

1. (Twice Amended) A compound having the formula I



I

and pharmaceutically acceptable salts and hydrates thereof, wherein:

A is selected from C1-alkyl optionally substituted with one to four halogen atoms, O(CH₂)₁₋₂, and S(CH₂)₁₋₂;

Ar is aryl or heteroaryl each optionally substituted with one to four groups independently selected from R8 selected from phenyl, 2-, 3-, 4-chlorophenyl, 2-, 3-, 4-bromophenyl, 2-, 3-, 4-fluorophenyl, 3,4-dichlorophenyl, 2,3-dichlorophenyl, 2,4-dichlorophenyl, 2,5-dichlorophenyl, 2,6-dichlorophenyl, 3,5-dichlorophenyl, 3-chloro-4-fluorophenyl, 2-chloro-4-fluorophenyl, 4-chloro-2-fluorophenyl, 2-cyanophenyl, 4-methylphenyl, 4-isopropylphenyl, 4-trifluoromethylphenyl, biphenyl, naphthyl, 3-methoxyphenyl, 3-carboxyphenyl, 2-carboxamidophenyl, 4-methoxyphenyl, 3-phenoxyphenyl, 4-(4-pyridyl)phenyl, 4-methylsulfonylphenyl, 3-dimethylaminophenyl, 5-tetrazolyl, 1-methyl-5-tetrazolyl, 2-methyl-5-tetrazolyl, 2-benzothienyl, 2-benzofuranyl, 2-indolyl, 2-quinolinyl, 7-quinolinyl, 2-benzothiazolyl, 2-benzimidazolyl, 1-benzotriazolyl, 2-furanyl, 3-furanyl, 2-imidazolyl, 5-imidazolyl, 5-isoxazolyl, 4-isoxazolyl, 4-isothiazolyl, 1,2,4-oxadiazol-5-yl, 2-oxazolyl, 4-oxazolyl, 4-pyrazolyl, 5-pyrazolyl, 2-pyridyl, 3-pyridyl, 2-pyrazinyl, 5-pyrimidinyl, 2-pyrrolyl, 4-thiazolyl, 1,2,4-thiadiazol-3-yl, 1,2,5-thiadiazol-4-yl, 1,2,3-thiadiazol-4-yl, 1,2,5-oxadiazol-4-yl, 1,2,3-oxadiazol-4-yl, 1,2,4-triazol-5-yl, 1,2,3-triazol-4-yl, 3-thienyl, 1,2,4-triazol-5-yl, pyrrolopyridine, furo[3,2-b]pyridin-2-yl, thieno[2,3-b]pyridin-2-yl, 5(H)-2-oxo-4-furanyl, 5(H)-2-oxo-5-furanyl, (1H,4H)-5-oxo-1,2,4-triazol-3-yl, 4-oxo-2-benzopyranyl;

Q is COOH,

one of X^1 , X^2 , or X^3 or X^4 is nitrogen and the others are independently selected from CH and $\text{C}-\text{Rg}$ and Rg is selected from 1) $\text{C}_1\text{-alkyl}$ optionally substituted with one to eight groups independently selected from aryl, heteroaryl, halogen, NR^aR^b , $\text{C}(\text{O})\text{R}^a$, $\text{C}(\text{OR}^a)\text{R}^a\text{R}^b$, SR^a and OR^a , wherein aryl, heteroaryl and alkyl are each optionally substituted with one to six groups independently selected from halogen, CF_3 , and COOH , or 2) $\text{S}(\text{O})_n\text{C}_1\text{-alkyl}$, wherein alkyl is optionally substituted with one to six substituents selected from halogen, aryl, heteroaryl, OH , and $\text{OC}(\text{O})\text{R}^a$;

X^2 is CH ;

X^4 is CH or $\text{C}-\text{Rg}$, where Rg is selected from 1) $\text{C}_1\text{-alkyl}$ optionally substituted with OR^a or 2) $\text{S}(\text{O})_n\text{C}_1\text{-alkyl}$;

Y^1 is S ;

Y^2 is selected from $(\text{CR}^d\text{R}^e)_m$ and $\text{CR}^d=\text{CRC}$;

R^1 is selected from H , CN , OR^a , $\text{S}(\text{O})_n\text{C}_1\text{-alkyl}$ and $\text{C}_1\text{-alkyl}$ optionally substituted with one to six groups independently selected from halogen, OR^a and $\text{S}(\text{O})_n\text{C}_1\text{-alkyl}$;

R^2 is selected from H and $\text{C}_1\text{-alkyl}$ optionally substituted with one to six halogen; or

R^3 is selected from H and $\text{C}_1\text{-alkyl}$ optionally substituted with one to six groups independently selected from OR^a and halogen;

R^a and R^b are independently selected from H and $\text{C}_1\text{-10alkyl}$, $\text{C}_2\text{-10alkenyl}$,

$\text{C}_2\text{-10alkynyl}$, Cy and $\text{Cy-C}_1\text{-10alkyl}$, wherein said alkyl, alkenyl, alkynyl and Cy are optionally substituted with one to six substituents independently selected from halogen, amino, carboxy, $\text{C}_1\text{-alkyl}$, $\text{C}_1\text{-alkoxy}$, aryl, heteroaryl, aryl- $\text{C}_1\text{-alkyl}$, hydroxy, CF_3 , $\text{OC}(\text{O})\text{C}_1\text{-alkyl}$, $\text{OC}(\text{O})\text{NR}^i\text{R}^j$, and aryl oxy; or

R^e is selected from $\text{C}_1\text{-alkyl}$ optionally substituted with one to six halogen, aryl and heteroaryl, wherein said aryl and heteroaryl are optionally substituted with one to three groups selected from halogen, $\text{OC}_1\text{-alkyl}$, $\text{O-haloC}_1\text{-alkyl}$, $\text{C}_1\text{-alkyl}$ and $\text{haloC}_1\text{-alkyl}$;

R^d and R^e are independently H , halogen, aryl, heteroaryl, $\text{C}_1\text{-alkyl}$ or $\text{haloC}_1\text{-alkyl}$;

R^f is selected from H , $\text{C}_1\text{-alkyl}$, $\text{haloC}_1\text{-alkyl}$, Cy , $\text{C}(\text{O})\text{C}_1\text{-alkyl}$, $\text{C}(\text{O})\text{haloC}_1\text{-alkyl}$, and $\text{C}(\text{O})\text{-Cy}$;

R^g is selected from

(1) — halogen,

(2) — CN ,

(3) — $\text{C}_1\text{-alkyl}$ optionally substituted with one to eight groups

independently selected from aryl, heteroaryl, halogen, NR^aR^b ,

$\text{C}(\text{O})\text{R}^a$, $\text{C}(\text{OR}^a)\text{R}^a\text{R}^b$, SR^a and OR^a , wherein aryl, heteroaryl and

alkyl are each optionally substituted with one to six groups independently selected from halogen, CF₃, and COOH;

(4) —C₂—alkenyl optionally substituted with one to six groups independently selected from halogen and OR^a;

(5) —Cy;

(6) —C(O)R^a;

(7) —C(O)OR^a;

(8) —CONR^aR^b;

(9) —OCONR^aR^b;

(10) —OC₁—alkyl, wherein alkyl is optionally substituted with one to six substituents selected from halogen, aryl, heteroaryl, OH and OC(O)R^a;

(11) —O—Cy;

(12) —S(O)_nC₁—alkyl, wherein alkyl is optionally substituted with one to six substituents selected from halogen, aryl, heteroaryl, OH, and OC(O)R^a;

(13) —S(O)_n—Cy;

(14) —NR^aS(O)_nR^b;

(15) —NR^aR^b;

(16) —NR^aC(O)R^b;

(17) —NR^aC(O)OR^b;

(18) —NR^aC(O)NR^aR^b;

(19) —S(O)_nNR^aR^b;

(20) —NO₂;

(21) —C₅—cycloalkenyl;

wherein Cy is optionally substituted with one to eight groups independently selected from halogen, C(O)R^a, OR^a, C₁—alkyl, aryl, heteroaryl and CF₃;

Rⁱ and R^j are independently selected from hydrogen, C₁—10alkyl, Cy and Cy—C₁—10alkyl; or Rⁱ and R^j together with the nitrogen atom to which they are attached form a ring of 5 to 7 members containing 0-2 additional heteroatoms independently selected from oxygen, sulfur and N—R^f;

Cy is selected from heterocyclyl, aryl, and heteroaryl;

m is 1 or 2; and

n is 0, 1 or 2.

2. (Original) A compound of Claim 1 wherein A-Q is CH₂CO₂H.

3. (Cancel)

4. (Previously Canceled)

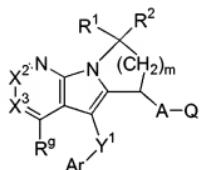
5. (Previously Canceled)

6. (Amended) A compound of Claim 1 wherein one of X¹, X² and X³ is nitrogen and the other is others are CH, X² is CH, and X⁴ is C-S(O)_n-C₁₋₆alkyl or C-C₁₋₆alkyl optionally substituted with OR^a.

7. (Cancel)

8. (Original) A compound of Claim 1 wherein Y² is selected from CH₂ and CH₂CH₂.

9. (Original) A compound of Claim 1 represented by the formula Ia:



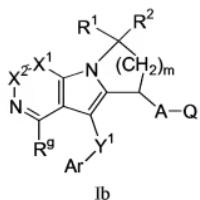
Ia

wherein X² and X³ are independently CH or C-Rg, A, Ar, Q, Y¹, R¹, R², m and Rg are as defined in Claim 1.

10. (Original) A compound of Claim 9 wherein X² and X³ are each CH, R¹ and R² are each H, and A-Q is CH₂CO₂H.

11. (Original) A compound of Claim 9 wherein Y¹-Ar is S-phenyl optionally substituted with 1 or 2 groups independently selected from halogen, C₁₋₆alkyl and trifluoromethyl.

12. (Original) A compound of Claim 1 represented by the formula Ib:

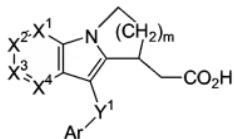


wherein X¹ and X² are independently CH or C-Rg, A, Ar, Q, Y¹, R¹, R², m and Rg are as defined in Claim 1.

13. (Original) A compound of Claim 12 wherein X¹ and X² are each CH, R¹ and R² are each H, and A-Q is CH₂CO₂H.

14. (Original) A compound of Claim 13 wherein Y¹-Ar is S-phenyl optionally substituted with 1 or 2 groups independently selected from halogen, C₁₋₆ alkyl and trifluoromethyl.

15. (Amended) A compound of Claim 1 represented by the formula Ic:



Ic

wherein one of X¹, X² and X³ is N and the others are each is CH, X⁴ is CH, X⁴ is CRg, m is 1 or 2, and Ar, Y¹ and m are as defined in Claim 1.

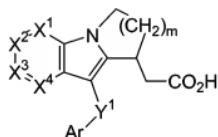
16. (Original) A compound of Claim 15 wherein Ar is phenyl optionally substituted with 1 or 2 groups independently selected from halogen, C₁₋₃alkyl and trifluoromethyl.

17. (Previously Canceled)

18. (Original) A compound of Claim 15 wherein X⁴ is selected from C-S(O)n-C₁₋₆alkyl and C-C₁₋₆alkyl optionally substituted with OR^a.

19. (Previously Amended) A compound of Claim 15 wherein Y¹-Ar is S-phenyl optionally substituted with 1 or 2 groups independently selected from halogen, C₁₋₆alkyl and trifluoromethyl; X¹ and X² are each CH, X³ is N, m is 1 or 2, and X⁴ is C-SO₂C₁₋₆alkyl or C-C₁₋₆alkyl.

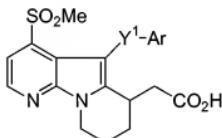
20. (Twice Amended) A compound of Claim 1 selected from:



X ¹	X ²	X ³	X ⁴	Ar	Y ¹	m
N	CH	CH	C(SO ₂ CH ₃)	4-Cl-Ph	S	2
N	CH	CH	C(SCH ₃)	4-Cl-Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	3,4-diCl-Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	4-Br-Ph	S	2
CH	CH	N	C(SO ₂ CH ₃)	3,4-diCl-Ph	S	1
CH	CH	N	C(SO ₂ CH ₃)	3,4-diCl-Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	4-CF ₃ -Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	2-Cl-4-F-Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	2-naphthyl	S	2
N	CH	CH	C(SO ₂ CH ₃)	2,3-diCl-Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	4-CH ₃ -Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	2,4-diCl-Ph	S	2
CH	CH	CH	C(SO ₂ CH ₃)	4-Cl-Ph	S	2
CH	C(CH ₃)	CH	C(SO ₂ CH ₃)	4-Cl-Ph	S	2
N	CH	C(CH ₃)	C(SO ₂ CH ₃)	4-Cl-Ph	S	2
CH	C(CH ₃)	N	C(SO ₂ CH ₃)	4-Cl-Ph	S	2
C(CH ₃)	CH	N	C(SO ₂ CH ₃)	4-Cl-Ph	S	2
N	CH	CH	C(CH(CH ₃) ₂)	4-F-Ph	S	2

X1	X2	X3	X4	Ar	Y1	m
N	CH	CH	C(CH(CH ₃) ₂)	4-Cl-Ph	S	2
N	CH	CH	C(CH(CH ₃) ₂)	2,4-diCl-Ph	S	2
N	CH	CH	C(CH(CH ₃) ₂)	4-Br-Ph	S	2
N	CH	CH	C(CH(CH ₃) ₂)	2-Cl-4-F-Ph	S	2
N	CH	CH	C(CH(CH ₃) ₂)	3,4-diCl-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	4-F-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	4-Cl-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	2,4-diCl-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	4-Br-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	2-Cl-4-F-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	3,4-diCl-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	4-F-Ph	S	1
CH	CH	N	C(CH(CH ₃) ₂)	4-Cl-Ph	S	1
CH	CH	N	C(CH(CH ₃) ₂)	2,4-diCl-Ph	S	1
CH	CH	N	C(CH(CH ₃) ₂)	4-Br-Ph	S	1
CH	CH	N	C(CH(CH ₃) ₂)	2-Cl-4-F-Ph	S	1
CH	CH	N	C(CH(CH ₃) ₂)	3,4-diCl-Ph	S	1
CH	N	CH	C(CH(CH ₃) ₂)	4-F-Ph	S	+
CH	N	CH	C(CH(CH ₃) ₂)	4-Cl-Ph	S	+
CH	N	CH	C(CH(CH ₃) ₂)	2,4-diCl-Ph	S	+
CH	N	CH	C(CH(CH ₃) ₂)	4-Br-Ph	S	+
CH	N	CH	C(CH(CH ₃) ₂)	2-Cl-4-F-Ph	S	+
CH	N	CH	C(CH(CH ₃) ₂)	3,4-diCl-Ph	S	+
CH	N	CH	C(CH(CH ₃) ₂)	4-F-Ph	S	2
CH	N	CH	C(CH(CH ₃) ₂)	4-Cl-Ph	S	2
CH	N	CH	C(CH(CH ₃) ₂)	2,4-diCl-Ph	S	2
CH	N	CH	C(CH(CH ₃) ₂)	4-Br-Ph	S	2
CH	N	CH	C(CH(CH ₃) ₂)	2-Cl-4-F-Ph	S	2
CH	N	CH	C(CH(CH ₃) ₂)	3,4-diCl-Ph	S	2
N	CH	CH	C(CH(OCH ₃) (CH ₂ CH ₃))	4-Cl-Ph	S	2
N	CH	CH	C(CH(OCH ₃) (CH ₂ CH ₃))	4-Cl-Ph	S	1
CH	N	CH	C(CH(OCH ₃) (CH ₂ CH ₃))	4-Cl-Ph	S	+

X1	X2	X3	X4	Ar	Y1	m
CH	N	CH	C(CH(OCH ₃)(CH ₂ CH ₃))	4-Cl-Ph	S	2
CH	CH	N	C(CH(OCH ₃)(CH ₂ CH ₃))	4-Cl-Ph	S	2
CH	CH	N	C(CH(OCH ₃)(CH ₂ CH ₃))	4-Cl-Ph	S	1
N	CH	CH	C(C(CH ₃) ₃)	4-Cl-Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	3,4-diCl-Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	4-Br-Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	4-CF ₃ -Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	2-Cl-4-F-Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	2-naphthyl	S	2
N	CH	CH	C(C(CH ₃) ₃)	2,3-diCl-Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	4-CH ₃ -Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	2,4-diCl-Ph	S	2



Ar	Y1
5-tetrazolyl	S
2-pyrrolyl	S
1,2,4-triazol-3-yl	S
1,2,3-triazol-4-yl	S
5-imidazolyl	S
4-pyrazolyl	S
5-pyrazolyl	S
(1H,4H)-5-oxo-1,2,4-triazol-3-yl	S
4-isothiazolyl	S
1,2,5-thiadiazol-5-yl	S
1,2,5-oxadiazol-5-yl	S

Ar	Y ¹
3-furanyl	S
1,2,3-thiadiazol-4-yl	S
1,2,3-oxadiazol-4-yl	S
4-isoxazolyl	S
3-thienyl	S
4-oxazolyl	S
4-thiazolyl	S
(5H)-2-oxo-5-furanyl	S
(5H)-2-oxo-4-furanyl	S
1,2,4-oxadiazol-5-yl	S
3-pyridyl	S
2-pyrazinyl	S
5-pyrimidinyl	S
2-indolyl	S
2-benzothienyl	S
2-benzofuranyl	S
4-oxo-benzopyran-2-yl	S
2-quinoliny	S
2-benzimidazolyl	S
2-benzoxazolyl	S
2-benzothiazolyl	S
1-benzotriazolyl	CH ₂ S
thieno[2,3-b]pyridin-2-yl	S

21. (Original) A pharmaceutical composition comprising a compound of Claim 1 and a pharmaceutically acceptable carrier.

22. (Cancel)

23. (Cancel)

24. (Cancel)

25. (Cancel)

26. (Cancel)

27. (Previously Canceled)

28. (Previously Canceled)

29. (Previously Canceled)

30. (Previously Canceled)